



0460 #4  
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**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

In re application of: Ewing et al.

Attorney Docket No.: CAMIP004

Application No.: 09/902,470

Examiner: UNASSIGNED

Filed: July 9, 2001

Group: UNASSIGNED

Title: ACCESSIBILITY CORRECTION FACTORS  
FOR ELECTRONIC MODELS OF  
CYTOCHROME P450 METABOLISM

**CERTIFICATE OF MAILING**

I hereby certify that this correspondence is being deposited with the United States Postal Service as First Class Mail to: Commissioner for Patents, Washington, DC, 20231 on July 27, 2001.

Signed: 

Leslie Russell

**INFORMATION DISCLOSURE STATEMENT  
37 CFR §§1.56 AND 1.97(b)**

Commissioner for Patents  
Washington, DC 20231

Dear Sir:


The references listed in the attached PTO Form 1449, copies of which are attached, may be material to examination of the above-identified patent application. Applicants submit these references in compliance with their duty of disclosure pursuant to 37 CFR §§1.56 and 1.97. The Examiner is requested to make these references of official record in this application.

This Information Disclosure Statement is not to be construed as a representation that a search has been made, that additional information material to the examination of this application does not exist, or that these references indeed constitute prior art.

This Information Disclosure Statement is: (i) filed within three (3) months of the filing date of the above-referenced application, (ii) believed to be filed before the mailing date of a first Office Action on the merits, or (iii) believed to be filed before the mailing of a first Office Action after the filing of a Request for Continued Examination under §1.114. Accordingly, it is believed that no fees are due in connection with the filing of this Information Disclosure Statement. However, if it is determined that any fees are due, the Commissioner is hereby authorized to charge such fees to Deposit Account 500388 (Order No. CAMIP004).

Respectfully submitted,

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#### U.S. Patent Documents

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	1A						

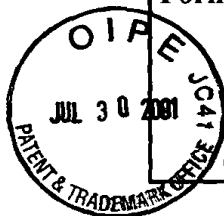
#### Foreign Patent or Published Foreign Patent Application

Examiner Initial	No.	Document No.	Publication Date	Country or Patent Office	Class	Sub-class	Translation	
							Yes	No
	1B	WO 95/18969	07/13/95	PCT				
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	1D	Bradford, M. M., et al., "A Rapid and Sensitive Method for the Quantitation of Microgram Quantities of Protein Utilizing the Principle of Protein-Dye Binding," ANAL. BIOCHEM., (1976) 72:248-54
	1E	Burka, L. T., et al., "Mechanism of Cytochrome P-450 Catalysis. Mechanism of N-Dealkylation and Amine Oxide Deoxygenation," J. AM. CHEM. SOC., (1985) 107:2549-51
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	1M	Gonzalez, F.J., et al., "Expression of Mammalian Cytochrome P450 Using Paccinia Virus," METHODS ENZYMOL., (1991) 206:85-92
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	3C	Heberger, K., "Linear Free Energy Relationships in Radical Reactions. II Hydrogen Abstraction From Substituted Toluenes by TERT-Butyl, TERT-Butoxyl and Tert-Butylperoxyl Radicals," J. PHYS. ORG. CHEM., (1994) 7:244-50
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	3K	Karki, S.B., et al., "On the Mechanism of Amine Oxidations by P450," Xenobiotica, (1995), 25(7):711-24
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	4C	Karki, S.B., et al., "Mechanism of Oxidative Amine Dealkylation of Substituted N,N-Dimethylanilines by Cytochrome P-450: Application of Isotope Effect Profiles," J. AM. CHEM. SOC., (1995) 117(13):3657-64
	4D	Kim, S.S.; et al., "Comparative Hammett Studies of Imidoyl, Benzylic, Aldehydic Hydrogens Transfer and Related Reaction by t-Butoxyl Radical," TETRAHEDRON LETT., (1985) 26(7): 891-4
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	4H	Korzekwa, K.R., et al., "Predicting the Cytochrome P450 Mediated Metabolism of Xenobiotics," PHARMACOGENETICS, (1993) 3:1-18
	4I	Lindsay Smith, J.R., et al., "Model Systems for Cytochrome P450 Dependent Mono-Oxygenases. Part 2. <sup>12</sup> Kinetic Isotope Effects for the Oxidative Demethylation of Anisole and [Me- <sup>2</sup> H <sub>3</sub> ] Anisole by Cytochrome P450 Dependent Mono-Oxygenases and Model Systems," J. CHEM. SOC. PERKIN TRANS. II, (1983) 5:621-8
	4J	Macdonald, T. L., et al., "Oxidation of Substituted N,N-Dimethylanilines by Cytochrome P-450: Estimation of the Effective Oxidation-Reduction Potential of Cytochrome P-450," (1989) BIOCHEMISTRY, 28:2071-7
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	5C	Manchester, J.I., et al., "A New Mechanistic Probe for Cytochrome P450: An Application of Isotope Effect Profiles," J. AM. CHEM. SOC., (1997) 119:5069-70
	5D	Nelson, D.R., et al., P450 Superfamily: Update on New Sequences, Gene Mapping, Accession Numbers and Nomenclature," PHARMACOGENETICS, (1996) 6:1-42
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	6C	Szklarz, G. D., et al., "Site-Directed Mutagenesis as a Tool for Molecular Modeling of Cytochrome P450 2B1," BIOCHEMISTRY, (1995) 34:14312-22
	6D	Tassaneeyakul, W., et al., "Human Cytochrome P450 Isoform Specificity in the Regioselective Metabolism of Toluene and o-, m- and p-Xylene," J. PHARMACOL. EXP. THER., (1996) 276(1):101-8
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	6L	Yin, H., et al., "Designing Safer Chemicals: Predicting the Rates of Metabolism of Halogenated Alkanes, " PROC. NATL. ACAD. SCI. USA, (1995) 92(24):11076-80
	6M	Zerner, Michael C., "Semiempirical Molecular Orbital Methods," REVIEWS IN COMPUTATIONAL CHEMISTRY II, Chapter 8, 313-365 (1991)
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	7C	International Search Report for PCT/LTS99/17713 dated 11 November 1999
	7D	Abstract No. XP-002122407, P84 to Johnson et al., "Automated Modeling Predicts Active Site Geometries Consistent with the Regiospecificity of P450s 2C3v and 2C5 for Progesterone Hydroxylation," FASEB Journal 11(9):P785 (1997)
	7E	Korzekwa and Gillette, "Overview: Theoretical Aspects of Isotope Effects on the Pattern of Metabolites Formed by Cytochrome P-450," Biological Reactive Intermediates IV, Witmer et al., Eds. Plenum Press, NY (1990)
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